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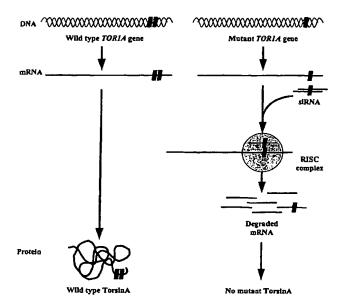
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(72) Inventors; and

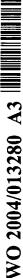
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- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW),

[Continued on next page]

(54) Title: ALLELE-SPECIFIC siRNA-MEDIATED GENE SILENCING



(57) Abstract: The present invention is directed to small interfering RNA molecules (siRNA) targeted against an allele of interest, and methods of using these siRNA molecules.



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Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

### Published:

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments

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### INTERNATIONAL SEARCH REPORT

Interestional Application is

PCT/US03/16887

A. CLASSIFICATION OF SUBJECT MATTER  IPC(7) : C12Q 1/68; C12P 19/34; C12N 15/63, 15/88; C07H 21/02, 21/04; A01N 43/04						
US CL : 435/6, 91.1, 91.31, 455, 458; 536/23.1, 24.5; 514/44						
According to International Patent Classification (IPC) or to both national classification and IPC						
B. FIELDS SEARCHED						
	locumentation searched (classification system follower 435/6, 91.1, 91.31, 455, 458; 536/23.1, 24.5; 514/44	d by classification symbols)	:			
Documenta	tion searched other than minimum documentation to the	ne extent that such documents are included	in the fields searched			
	lata base consulted during the international search (na gg, STIC sequence search	me of data base and, where practicable, sea	urch terms used)			
C. DO	CUMENTS CONSIDERED TO BE RELEVANT					
Category *	Citation of document, with indication, where	<del></del>	Relevant to claim No.			
Y	US 6,387,616 (ORZELIUS et al.) 14 May 2002 (at ID NO. 24).	ostract, col. 1, 7, 12-14, 17, 51-52, SEQ	1-17, 25-27			
Y	FIRE, A. et al., Potent and specific genetic interfer elegans, Nature, Vol. 391, pages 806-811, Feb. 19, and Tablé 3 on p. 809).					
Y	US 5,902,880 (THOMPSON) 11 May 1999 (col. 6	1-17, 25-27				
Y	US 2002/0086356 (TUSCHL et al.) 4 July 2002 (p.	3-8				
Further	er documents are listed in the continuation of Box C.	See patent family annex.				
* Special categories of cited documents:  "A" document defining the general state of the art which is not considered to be of		"I" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention				
"B" carliera	ier application or patent published on or after the international filing date considered novel or cann when the document is tak		imed invention cannot be d to involve an inventive step			
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"O" document referring to an oral disclosure, use, exhibition or other means		to a person skilled in the art	_			
"P" document published prior to the international filing date but fater than the priority date claimed		"&" document member of the same patent for	•			
Date of the actual completion of the international search		Date of mailing of the international cearch report  31 00 2005  Authorized officer  Jane Zara				
14 September 2005 (14.09.2005)  Name and mailing address of the ISA/US		Authorized office	, ,			
Mai	1 Stop PCT, Attn: ISA/US	Januar Ohn	and 1			
Con	mnissioner of Patents D. Box 1450	Jane Zara	•			
Ala	szandria, Virginia 22313-1450	Telephone No. (701) 208-0196				
	o. (703) 305-3230 A/210 (second sheet) (July 1998)	L				
	and from minds (and 1990)					

### INTERNATIONAL SEARCH REPORT

Interestimal audication Action PCT/US03/16887

Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)					
This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:					
Claim Nos.:  because they relate to subject matter not required to be searched by this Authority, namely.  Claims 18-24 and 41-60 were not searchable because they were multiply dependent claims.					
Claim Nos.: 18-24 and 41-60  because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically.					
Claim Nos.:     because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).					
Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)					
This International Searching Authority found multiple inventions in this international application, as follows: Please See Continuation Sheet					
1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.					
<ol> <li>As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.</li> </ol>					
3. As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:					
4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is					
restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-17,25-27 and 61-81					
Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.					

Form PCT/ISA/210 (continuation of first sheet(1)) (July 1998)

# PATENT COOPERATION TREATY

	From t	From the INTERNATIONAL BUREAU				
PCT	To:	То:				
NOTIFICATION OF THE RECORDING	VIKS	VIKSNINS, Ann, S.				
OF A CHANGE		Fish & Richardson P.C., P.A.				
(PCT Rule 92bis.1 and		60 South Sixth Street				
Administrative Instructions, Section 422)		Suite 3300 Minneapolis, MN 55402				
		United States of America				
Date of mailing (day/month/year)	ļļ					
02 February 2005 (02.02.2005)	<u> </u>					
Applicant's or agent's file reference		IMPORTANT NOTIFICATION				
17023-045WO1	1	INFORTANT NOTIFICATION				
International application No.	Internation	International filing date (day/month/year)				
PCT/US2003/016887	26 N	26 May 2003 (26.05.2003)				
1. The following indications appeared on record concerning:    X   the applicant   X   the inventor	the ager	the samme	n representative			
X the applicant X the inventor	The ager		·			
Name and Address		State of Nationality	State of Residence			
HARPER, Scott Department of Neurology		US Telephone No.	US			
University of Iowa College of		relephone ivo.				
Medicine 2007 RCP		Facsimile No.				
lowa City, IA 52242 United States of America						
Sinisa States St. Amorios		Teleprinter No.				
			:			
2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:						
the person the name X the add	dress	the nationality	the residence			
Name and Address		State of Nationality	State of Residence			
HARPER, Scott		US	US			
1 Warwick Circle lowa City, IA 52240-2847 United States of America  EPO -D	C 1	Telephone No.				
United States of America	ا الحديث					
1 6. 02. 20	nns	Facsimile No.				
. <b> </b>	JUJ	Teleprinter No.				
(117)	:	relephinter (vo.				
3. Further observations, if necessary:						
4. A copy of this notification has been sent to:						
X the receiving Office	. Г	X the designated Offices concerned				
X the International Searching Authority	L F	the elected Offices concerned				
the International Preliminary Examining Authority	Ĺ	other:				
the international Freignmary Examining Authority						
The International Bureau of WIPO		Authorized officer				
34, chemin des Colombettes	j	Anne KARKA	CHI (Fax 338.87.20)			
1211 Geneva 20, Switzerland						
Facsimile No. (41-22) 338.87.20		Telephone No. (41-22) 338 9656				